SEQUENCE LISTING



GENERAL INFORMATION:

- (i) APPLICANT: Martin Roland Jensen Soren Mouritsen Henrik Elsner Iben Dalum
- (ii) TITLE OF INVENTION: Modified human TNF-alpha molecules, DNA encoding them, and vaccines containing said modified TNF-alpha or DNA
- (iii) NUMBER OF SEQUENCES: 44
- (iv) CORRESPONDENCE ADDRESS:
 - (A) ADDRESSEE: JACOBSON HOLMAN, PLLC
 - (B) STREET: 400 Seventh St., N.W.
 - (C) CITY: Washington
 - (D) STATE: DC
 - (E) COUNTRY: USA
 - (F) POSTAL CODE: 20004
- (v) COMPUTER READABLE FORM:
 - (A) MEDIUM TYPE: Floppy disk
 - (B) COMPUTER: IBM PC compatible
 - (C) OPERATING SYSTEM: PC-DOS/MS-DOS
 - (D) SOFTWARE: PatentIn Release #1.0, Version #1.30 (EPO)
- (vi) CURRENT APPLICATION DATA:
 - (A) APPLICATION NUMBER: 09/060,294
 - (B) FILING DATE: 15-APR-1998
- (vii) PRIOR APPLICATION DATA:
 - (A) APPLICATION NUMBER: 60/044,187
 - (B) FILING DATE: 24-APR-1997
- (viii) ATTORNEY/AGENT INFORMATION:
 - (A) NAME: William E. Player
 - (B) REGISTRATION NUMBER: 31,409
 - (C) REFERENCE/DOCKET NUMBER: P60953US1
- (ix) TELECOMMUNICATION INFORMATION:
 - (A) TELEPHONE: (202) 638-6666
 - (B) TELEFAX: (202) 393-5350
- (2) INFORMATION FOR SEQ ID NO: 1:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: cDNA
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO

(vi) ORIGINAL SOURCE:

(A) ORGANISM: Homo sapiens

(ix) FEATURE:

- (A) NAME/KEY: CDS
- (B) LOCATION:1..474
- (C) IDENTIFICATION METHOD: experimental
- (D) OTHER INFORMATION:/codon_start= 1
 /function= "Antigen"
 /product= "TNF-alpha analog"
 /evidence= EXPERIMENTAL
 /gene= "tnfP2-1"
 /standard_name= "TNF2-1"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

		TCT Ser 5						48
		GGT Gly						96
		CTC Leu						144
	 	 TCA Ser	 					192
		GGC Gly						240
	 	 GCC Ala 85						288
		CCC Pro						336
		GAG Glu						384
-		CTC Leu						432
	 	 GGG Gly						474

(2) INFORMATION FOR SEQ ID NO: 2:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:
- Met Val Arg Ser Ser Ser Arg Thr Pro Ser Gln Tyr Ile Lys Ala Asn 1 5 10 15
- Ser Lys Phe Ile Gly Ile Thr Glu Leu Gln Leu Gln Trp Leu Asn Arg
 20 25 30
- Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln 35 40 45
- Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 55 60
- Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80
- Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95
- Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110
- Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu
 115 120 125
- Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140
- Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155
- (2) INFORMATION FOR SEQ ID NO: 3:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION: 1..474

(D) OTHER INFORMATION:/codon_start= 1
 /function= "Antigen"
 /product= "TNF-alpha analog"
 /gene= "tnfP2-3"
 /standard_name= "TNF2-3"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3:

					GAC Asp 170			48
					CTC Leu			96
					GAG Glu			144
					ATC Ile			192
					ATC Ile			240
-					AAG Lys 250			288
					CCA Pro			336
					GGG Gly			384
					CGG Arg			432
					ATC Ile			474

(2) INFORMATION FOR SEQ ID NO: 4:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

- Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His

 1 10 15
- Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg
 20 25 30
- Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln
 35 40 45
- Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 60
- Phe Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly Ile Thr Glu Leu 65 70 75 80
- Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95
- Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110
- Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu 115 120 125
- Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140
- Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155
- (2) INFORMATION FOR SEQ ID NO: 5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION: 1..474
 - (D) OTHER INFORMATION:/codon_start= 1
 /function= "Antigen"
 /product= "TNF-alpha analog"
 /gene= "tnfP2-4"
 /standard name= "TNF2-4"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

Met 160	Val	Arg	Ser	Ser	Ser 165	Arg	Thr	Pro	Ser	Asp 170	Lys	Pro	Val	Ala	His 175	
					CAA Gln											96
					CTG Leu											144
					GAG Glu											192
					TGC Cys											240
					GTC Val 245											288
					TGC Cys											336
					AAG Lys											384
					AGC Ser											432
		GAG			CAG Gln		TAC					GCC				474

(2) INFORMATION FOR SEQ ID NO: 6:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His 1 5 10 15

Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg 20 25 30

Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln 35 40 45

Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu

50		55	60

Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80

Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95

Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110

Lys Pro Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly Ile Thr Glu 115 120 125

Leu Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140

Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155

(2) INFORMATION FOR SEO ID NO: 7:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO
- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION: 1..474
 - (D) OTHER INFORMATION:/function= "Antigen"
 /product= "TNF-alpha analog"
 /gene= "tnfP2-5"
 /standard_name= "TNF2-5"
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 7:

ATG	GTC	AGA	TCA	TCT	TCT	CGA	ACC	CCG	AGT	GAC	AAG	CCT	GTA	GCC	CAT	48
Met	Val	Arg	Ser	Ser	Ser	Arg	Thr	Pro	Ser	Asp	Lys	Pro	Val	Ala	His	
160					165					170					175	

GTT GTA GCA AAC CCT CAA GCT GAG GGG CAG CTC CAG TGG CTG AAC CGC 96
Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg
180 185 190

CGG GCC AAT GCC CTC CTG GCC AAT GGC GTG GAG CTG AGA GAT AAC CAG
Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln
195 200 205

		GAG Glu						192
		TGC Cys						240
		GTC Val 245						288
		TGC Cys						336
		CCC Pro						384
		TAC Tyr						432
		CAG Gln						474

(2) INFORMATION FOR SEQ ID NO: 8:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 8:

Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His 1 5 10 15

Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg
20 25 30

Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln 35 40 45

Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 55 60

Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80

Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser

Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110

Lys	Pro	Trp 115	Tyr	Glu	Pro	Ile	Tyr 120	Leu	Gly	Gly	Val	Phe 125	Gln	Leu	Glu	
Lys	Gly 130	Asp	Arg	Gln	Tyr	Ile 135	Lys	Ala	Asn	Ser	Lys 140	Phe	Ile	Gly	Ile	
Thr 145	Glu	Leu	Ser	Gly	Gln 150	Val	Tyr	Phe	Gly	Ile 155	Ile	Ala	Leu			
(2)	INFO	ORMA	rion	FOR	SEQ	ID 1	10: 9	9:								
	(i)	(I (C	A) Li 3) Ti C) Si	CE CHENGTH YPE: TRANI	H: 47 nucl	74 ba leic ESS:	ase p acid doul	pairs d	5							
	(ii)	MOI	LECUI	LE T	YPE:	DNA	(gei	nomi	c)							
	(iii)	HYI	РОТНІ	ETICA	AL: 1	40										
	(iv)	ANT	ri-si	ENSE	: NO											
	(vi)			AL SO			o sap	piens	3							
	(ix)	(<i>1</i> (1	3) L(AME/I OCAT: THER /fi /pi /ge	ION: 1 INFO uncti coduce ene=	147	TION "Ant TNF- EP2-	tiger -alph 7"	n" na ar	nalog						
	(xi)	SE	QUENC	CE DE	ESCRI	IPTIC	ON: S	SEQ :	ID NO	0: 9	:					
	GTC Val															48
	GTA Val															96
	GCC Ala															144
	GTG Val															192
	AAG Lys 225															240
GCT	AAC	TCC	AAA	TTC	ATC	GGC	ATC	ACC	GAA	CTG	GTT	AAC	CTC	CTC	TCT	288

Ala 240	Asn	Ser	Lys	Phe	Ile 245	Gly	Ile	Thr	Glu	Leu 250	Val	Asn	Leu	Leu	Ser 255	
											GAG Glu					336
											GTC Val					384
											CCC Pro					432
											ATT Ile 315					474
(2)	INFO	RMAT	CION	FOR	SEQ	ID N	10: 1	10:								
	!	(E	SEQUE A) LE B) TY D) TO	ENGTH PE:	I: 15 amin	8 am 10 ac	nino cid									
	(ii)	MOI	ECUL	ьЕ ТУ	PE:	prot	ein									
	(xi)	SEÇ	UENC	CE DE	SCRI	PTIC	N: S	SEQ 1	D NO): 10):					
Met 1	Val	Arg	Ser	Ser 5	Ser	Arg	Thr	Pro	Ser 10	Asp	Lys	Pro	Val	Ala 15	His	
Val	Val	Ala	Asn 20	Pro	Gln	Ala	Glu	Gly 25	Gln	Leu	Gln	Trp	Leu 30	Asn	Arg	
Arg	Ala	Asn 35	Ala	Leu	Leu	Ala	Asn 40	Gly	Val	Glu	Leu	Arg 45	Asp	Asn	Gln	
Leu	Val 50	Val	Pro	Ser	Glu	Gly 55	Leu	Tyr	Leu	Ile	Tyr 60	Ser	Gln	Val	Leu	
Phe 65	Lys	Gly	Gln	Gly	Cys 70	Pro	Ser	Thr	His	Val 75	Leu	Gln	Tyr	Ile	Lys 80	
Ala	Asn	Ser	Lys	Phe 85	Ile	Gly	Ile	Thr	Glu 90	Leu	Val	Asn	Leu	Leu 95	Ser	
Ala	Ile	Lys	Ser 100	Pro	Cys	Gln	Arg	Glu 105	Thr	Pro	Glu	Gly	Ala 110	Glu	Ala	
Lys	Pro	Trp 115	Tyr	Glu	Pro	Ile	Tyr 120	Leu	Gly	Gly	Val	Phe 125	Gln	Leu	Glu	
Lys	Gly 130	Asp	Arg	Leu	Ser	Ala 135	Glu	Ile	Asn	Arg	Pro 140	Asp	Tyr	Leu	Asp	
Phe 145	Ala	Glu	Ser	Gly	Gln 150	Val	Tyr	Phe	Gly	Ile 155	Ile	Ala	Leu			

(A) LENGTH: 474 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: double (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..474 (D) OTHER INFORMATION:/codon start= 1 /function= "Antigen" /product= "TNF-alpha analog" /gene= "tnfP30-1" /standard name= "TNF30-1" (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 11: ATG GTC AGA TCA TCT TCT CGA ACC CCG AGT TTC AAC AAT TTT ACC GTA 48 Met Val Arg Ser Ser Ser Arg Thr Pro Ser Phe Asn Asn Phe Thr Val 160 165 170 AGC TTT TGG CTC CGT GTA CCT AAG GTG TCG GCC TCG CAC CTG GAG CGC 96 Ser Phe Trp Leu Arg Val Pro Lys Val Ser Ala Ser His Leu Glu Arg 180 185 CGG GCC AAT GCC CTC CTG GCC AAT GGC GTG GAG CTG AGA GAT AAC CAG 144 Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln 195 200 CTG GTG GTG CCA TCA GAG GGC CTG TAC CTC ATC TAC TCC CAG GTC CTC 192 Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 215 210 TTC AAG GGC CAA GGC TGC CCC TCC ACC CAT GTG CTC CTC ACC CAC ACC 240 Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 225 230 ATC AGC CGC ATC GCC GTC TCC TAC CAG ACC AAG GTC AAC CTC CTC TCT 288 Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 245 250 GCC ATC AAG AGC CCC TGC CAG AGG GAG ACC CCA GAG GGG GCT GAG GCC 336 Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 260 265 AAG CCC TGG TAT GAG CCC ATC TAT CTG GGA GGG GTC TTC CAG CTG GAG 384 Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu

(2) INFORMATION FOR SEQ ID NO: 11:

(i) SEQUENCE CHARACTERISTICS:

275 280 285

AAG GGT GAC CGA CTC AGC GCT GAG ATC AAT CGG CCC GAC TAT CTC GAC
Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp
290
295
300

TTT GCC GAG TCT GGG CAG GTC TAC TTT GGG ATC ATT GCC CTC

Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu

305

315

(2) INFORMATION FOR SEQ ID NO: 12:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 12:

Met Val Arg Ser Ser Ser Arg Thr Pro Ser Phe Asn Asn Phe Thr Val

1 1 1 15

Ser Phe Trp Leu Arg Val Pro Lys Val Ser Ala Ser His Leu Glu Arg
20 25 30

Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln
35 40 45

Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 55 60

Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80

Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95

Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110

Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu 115 120 125

Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 140

Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155

(2) INFORMATION FOR SEQ ID NO: 13:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear

(iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: CDS (B) LOCATION: 1..474 (D) OTHER INFORMATION:/codon start= 1 /function= "Antigen" /product= "TNF-alpha analog" /gene= "tnfP30-2" /standard name= "TNF30-2" (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 13: ATG GTC AGA TCA TCT TCT CGA ACC CCG AGT GAC AAG CCT GTA GCC CAT 48 Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His 170 165 GTT GTA GCA AAC CCT CAA GCT GAG GGG CAG CTC CAG TGG CTG AAC CGC 96 Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg 185 180 CGG GCC AAT GCC CTC CTG GCC AAT TTC AAC AAC TTC ACA GTT AGC TTC 144 Arg Ala Asn Ala Leu Leu Ala Asn Phe Asn Asn Phe Thr Val Ser Phe 195 200 TGG TTG AGG GTA CCA AAG GTC TCG GCC AGC CAC CTC GAG CAG GTC CTC 192 Trp Leu Arg Val Pro Lys Val Ser Ala Ser His Leu Glu Gln Val Leu 210 215 TTC AAG GGC CAA GGC TGC CCC TCC ACC CAT GTG CTC CTC ACC CAC ACC 240 Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 225 230 235 ATC AGC CGC ATC GCC GTC TCC TAC CAG ACC AAG GTC AAC CTC CTC TCT 288 Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 250 245 GCC ATC AAG AGC CCC TGC CAG AGG GAG ACC CCA GAG GGG GCT GAG GCC 336 Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 260 265 AAG CCC TGG TAT GAG CCC ATC TAT CTG GGA GGG GTC TTC CAG CTG GAG 384 Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu 275 280 AAG GGT GAC CGA CTC AGC GCT GAG ATC AAT CGG CCC GAC TAT CTC GAC 432 Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 290 295 TTT GCC GAG TCT GGG CAG GTC TAC TTT GGG ATC ATT GCC CTC 474 Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 305 310 315

(ii) MOLECULE TYPE: DNA (genomic)

- (2) INFORMATION FOR SEQ ID NO: 14:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: protein
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14:
- Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His

 1 10 15
- Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg 20 25 30
- Arg Ala Asn Ala Leu Leu Ala Asn Phe Asn Asn Phe Thr Val Ser Phe 35 40 45
- Trp Leu Arg Val Pro Lys Val Ser Ala Ser His Leu Glu Gln Val Leu 50 60
- Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80
- Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95
- Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110
- Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu
 115 120 125
- Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140
- Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155
- (2) INFORMATION FOR SEQ ID NO: 15:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens

(ix) FEATURE:

- (A) NAME/KEY: CDS
- (B) LOCATION:1..474
- (D) OTHER INFORMATION:/codon_start= 1 /function= "Antigen" /product= "TNF-alpha analog"

/gene= "tnfP30-3"

/standard_name= "TNF30-3"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15:

			CGA Arg					_	48
			GCT Ala						96
			GCC Ala						144
			GGC Gly						192
			TCC Ser 230						240
			TCC Ser						288
			CAG Gln						336
			ATC Ile						384
			GCT Ala						432
			GTC Val 310						474

(2) INFORMATION FOR SEQ ID NO: 16:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids (B) TYPE: amino acid

 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 16:
- Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His 1 5 10 15
- Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg
 20 25 30
- Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln 35 40 45
- Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 60
- Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg Val Pro Lys Val Ser 65 70 75 80
- Ala Ser His Leu Glu Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95
- Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala 100 105 110
- Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu 115 120 125
- Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140
- Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155
- (2) INFORMATION FOR SEQ ID NO: 17:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION:1..474
 - (D) OTHER INFORMATION:/function= "Antigen"
 /product= "TNF-alpha analog"
 /gene= "tnfP30-4"
 /standard name= "TNF30-4"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

				Arg					48
				GCT Ala					96
				GCC Ala					144
				GGC Gly					192
_	-	-		CCC Pro 230					240
				TCC Ser					288
				CAG Gln					336
				GTC Val					384
				GCT Ala					432
		Glu		GTC Val 310					474

(2) INFORMATION FOR SEQ ID NO: 18:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 158 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

Met Val Arg Ser Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His 1 5 10 15

Val Val Ala Asn Pro Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg 20 25 30

Arg Ala Asn Ala Leu Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln

35	40	45
	- •	

Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu 50 60

Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr 65 70 75 80

Ile Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser 85 90 95

Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Phe Asn Asn Phe Thr
100 105 110

Val Ser Phe Trp Leu Arg Val Pro Lys Val Ser Ala Ser His Leu Glu 115 120 125

Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp 130 135 140

Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu 145 150 155

(2) INFORMATION FOR SEQ ID NO: 19:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 474 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO
- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION: 1..474
 - (D) OTHER INFORMATION:/codon_start= 1
 /function= "Antigen"
 /product= "TNF-alpha analog"
 /gene= "tnfP30-5"
 /standard name= "TNF30-5"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 19:

 GTC Val								48
 GTA Val								96

Arg	Ala	Asn	Ala 195	Leu	Leu	Ala	Asn	Gly 200	Val	Glu	Leu	Arg	Asp 205	Asn	Gln	
					GAG Glu											192
					TGC Cys											240
					GTC Val 245											288
					TGC Cys											336
					CCC Pro											384
					AAC Asn											432
					TCG Ser											474
(2)	INFO	ORMAT	CION	FOR	SEQ	ID N	10: 2	20:								
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 158 amino acids (B) TYPE: amino acid (D) TOPOLOGY: linear																
(ii) MOLECULE TYPE: protein																
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 20:																
Met 1	Val	Arg	Ser	Ser 5	Ser	Arg	Thr	Pro	Ser 10	Asp	Lys	Pro	Val	Ala 15	His	
Val																
	Val	Ala	Asn 20	Pro	Gln	Ala	Glu	Gly 25	Gln	Leu	Gln	Trp	Leu 30	Asn	Arg	
Arg			20		Gln Leu			25					30		_	
_	Ala	Asn 35	20 Ala	Leu		Ala	Asn 40	25 Gly	Val	Glu	Leu	Arg 45	30 Asp	Asn	Gln	
Leu	Ala Val 50	Asn 35 Val	20 Ala Pro	Leu Ser	Leu	Ala Gly 55	Asn 40 Leu	25 Gly Tyr	Val Leu	Glu Ile	Leu Tyr 60	Arg 45 Ser	30 Asp Gln	Asn Val	Gln Leu	

```
Ala Ile Lys Ser Pro Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala
Lys Pro Trp Tyr Glu Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu
        115
Lys Gly Asp Arg Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg Val
                        135
Pro Lys Val Ser Ala Ser His Leu Glu Gly Ile Ile Ala Leu
                    150
145
(2) INFORMATION FOR SEQ ID NO: 21:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 24 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (ix) FEATURE:
          (A) NAME/KEY: misc_feature
          (B) LOCATION:1..24
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha"
                 /product= "Primer binding to TNF-alpha gene"
                 /evidence= EXPERIMENTAL
                 /standard_name= "TNF-alpha Primer I"
                 /label= Primer1
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 21:
                                                                         24
GACAAGCCCA TGGTCAGATC ATCT
```

(2) INFORMATION FOR SEQ ID NO: 22:

(i) SEQUENCE CHARACTERISTICS:

- (A) LENGTH: 30 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO
- (ix) FEATURE:
 - (A) NAME/KEY: misc_feature

```
/standard name= "TNF-alpha Primer II"
                 /label= Primer2
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 22:
                                                                         30
TCTCTAGAGG GCAATGATCC CAAAGTAGAC
(2) INFORMATION FOR SEQ ID NO: 23:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 21 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (ix) FEATURE:
          (A) NAME/KEY: misc_feature
          (B) LOCATION:1..21
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha"
                 /product= "Primer binding to TNF-alpha gene"
                 /evidence= EXPERIMENTAL
                 /standard name= "TNF-alpha Primer III"
                 /label= Primer3
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 23:
CCCAAAGTAG ACCTGCCCAG A
                                                                         21
(2) INFORMATION FOR SEQ ID NO: 24:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 69 base pairs
          (B) TYPE: nucleic acid
```

(B) LOCATION:1..30

(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(A) ORGANISM: Homo sapiens

(ii) MOLECULE TYPE: DNA (genomic)

(iii) HYPOTHETICAL: NO

(iv) ANTI-SENSE: NO

(vi) ORIGINAL SOURCE:

(C) IDENTIFICATION METHOD: experimental

of DNA encoding TNF-alpha"

/evidence= EXPERIMENTAL

(D) OTHER INFORMATION:/function= "Primer for PCR cloning

/product= "Primer binding to TNF-alpha gene"

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Section 1

```
(ix) FEATURE:
          (A) NAME/KEY: insertion seq
          (B) LOCATION: 7..51
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut2-1""
                 /label= mut2-1
                 /note= "Primer "mut2-1" is a synthetically synthesised
                 69-mer oligonucleotide comprising DNA encoding the human
                 T cell epitope P2 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 24:
ACCCCGAGTC AGTACATTAA AGCCAATTCT AAATTCATCG GTATAACTGA GCTGCAGCTC
                                                                        60
CAGTGGCTG
                                                                        69
(2) INFORMATION FOR SEQ ID NO: 25:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 73 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (vi) ORIGINAL SOURCE:
          (A) ORGANISM: Homo sapiens
    (ix) FEATURE:
          (A) NAME/KEY: insertion_seq
          (B) LOCATION: 15...59
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut2-3""
                 /label= mut2-3
                 /note= "Primer "mut2-3" is a synthetically synthesised
                 73-mer oligonucleotide comprising DNA encoding the human
                 T cell epitope P2 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 25:
```

73

CCCAGGTCCT CTTCCAGTAC ATAAAGGCCA ACTCCAAGTT TATCGGCATC ACCGAGCTCA

TCAGCCGCAT CGC

(2) INFORMATION FOR SEQ ID NO: 26: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 75 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens (ix) FEATURE: (A) NAME/KEY: insertion seq (B) LOCATION: 12..56 (C) IDENTIFICATION METHOD: experimental (D) OTHER INFORMATION:/function= "Primer for PCR cloning of DNA encoding TNF-alpha analog" /evidence= EXPERIMENTAL /organism= "Homo sapiens" /standard_name= "Primer "mut2-4"" /label= mut2-4 /note= "Primer "mut2-4" is a synthetically synthesised 75-mer oligonucleotide comprising DNA encoding the human T cell epitope P2 between stretches of DNA homologous to stretches of the human TNF-alpha gene" (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 26: AGTCGGTCAC CGAGCTCCGT GATGCCGATG AATTTCGAAT TGGCCTTGAT ATACTGGGGC 60 TTGGCCTCAG CCCCC 75 (2) INFORMATION FOR SEQ ID NO: 27: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 75 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: DNA (genomic) (iii) HYPOTHETICAL: NO (iv) ANTI-SENSE: NO (vi) ORIGINAL SOURCE: (A) ORGANISM: Homo sapiens

(ix) FEATURE:

(A) NAME/KEY: insertion_seq

(B) LOCATION:8..52

(C) IDENTIFICATION METHOD: experimental

```
(D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut2-5""
                 /label= mut2-5
                 /note= "Primer "mut2-5" is a synthetically synthesised
                 75-mer oligonucleotide comprising DNA encoding the human
                 T cell epitope P2 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 27:
GAAGGGTGAC CGACAGTACA TTAAGGCCAA TTCGAAGTTC ATTGGCATCA CTGAGCTGTC
                                                                        60
TGGGCAGGTC TACTT
                                                                        75
(2) INFORMATION FOR SEQ ID NO: 28:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 80 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (vi) ORIGINAL SOURCE:
          (A) ORGANISM: Homo sapiens
    (ix) FEATURE:
          (A) NAME/KEY: insertion seq
          (B) LOCATION: 14..58
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut2-7""
                 /label= mut2-7
                 /note= "Primer "mut2-7" is a synthetically synthesised
                 80-mer oligonucleotide comprising DNA encoding the human
                 T cell epitope P2 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 28:
CACCCATGTG CTCCAGTACA TCAAAGCTAA CTCCAAATTC ATCGGCATCA CCGAACTGGT
                                                                        60
TAACCTCCTC TCTGCCATCA
                                                                        80
```

(2) INFORMATION FOR SEQ ID NO: 29:

```
(i) SEQUENCE CHARACTERISTICS:
```

- (A) LENGTH: 96 base pairs
- (B) TYPE: nucleic acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: DNA (genomic)
- (iii) HYPOTHETICAL: NO
- (iv) ANTI-SENSE: NO
- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
 - (A) NAME/KEY: insertion seq
 - (B) LOCATION: 10..72
 - (C) IDENTIFICATION METHOD: experimental
 - (D) OTHER INFORMATION:/function= "Primer for PCR cloning of DNA encoding TNF-alpha analog" /evidence= EXPERIMENTAL /organism= "Homo sapiens" /standard_name= "Primer "mut30-1"" /label= mut30-1 /note= "Primer "mut30-1" is a synthetically synthesised 96-mer oligonucleotide comprising DNA encoding the human T cell epitope P30 between stretches of DNA homologous to stretches of the human TNF-alpha gene"
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 29:

ACCCCGAGTT TCAACAATTT TACCGTAAGC TTTTGGCTCC GTGTACCTAA GGTGTCGGCC

TCGCACCTGG AGCGCCGGGC CAATGCCCTC CTGGCC

96

60

- (2) INFORMATION FOR SEQ ID NO: 30:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 100 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: DNA (genomic)
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: insertion_seq
 - (B) LOCATION: 12..74
 - (C) IDENTIFICATION METHOD: experimental
 - (D) OTHER INFORMATION:/function= "Primer for PCR cloning of DNA encoding TNF-alpha analog" /evidence= EXPERIMENTAL

```
/standard name= "Primer "mut30-2""
                 /label= mut30-2
                 /note= "Primer "mut30-2" is a synthetically synthesised
                 100-mer oligonucleotide comprising DNA encoding human T
                 cell epitope P30 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 30:
TCCTGGCCAA TTTCAACAAC TTCACAGTTA GCTTCTGGTT GAGGGTACCA AAGGTCTCGG
                                                                        60
CCAGCCACCT CGAGCAGGTC CTCTTCAAGG GCCAAGGCTG
                                                                       100
(2) INFORMATION FOR SEQ ID NO: 31:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 100 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (vi) ORIGINAL SOURCE:
          (A) ORGANISM: Homo sapiens
    (ix) FEATURE:
          (A) NAME/KEY: insertion seq
          (B) LOCATION: 12..74
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION: /function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut30-3""
                 /label= mut30-3
                 /note= "Primer "mut30-3" is a synthetically synthesised
                 100-mer oligonucleotide comprising DNA encoding human T
                 cell epitope P30 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 31:
CCCAGGTCCT CTTCAACAAC TTTACCGTCT CCTTCTGGCT TCGGGTACCC AAGGTCAGCG
                                                                        60
CTAGCCACCT CGAGGTCTCC TACCAGACCA AGGTCAACCT
                                                                       100
(2) INFORMATION FOR SEQ ID NO: 32:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 100 base pairs
          (B) TYPE: nucleic acid
```

/organism= "Homo sapiens"

(C) STRANDEDNESS: single

```
(D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (vi) ORIGINAL SOURCE:
          (A) ORGANISM: Homo sapiens
    (ix) FEATURE:
          (A) NAME/KEY: insertion seq
          (B) LOCATION: 15..77
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard_name= "Primer "mut30-4""
                 /label= mut30-4
                 /note= "Primer "mut30-4" is a synthetically synthesised
                 100-mer oligonucleotide comprising DNA encoding human T
                 cell epitope P30 between stretches of DNA homologous to
                 stretches of the human TNF-alpha gene"
    (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 32:
AGTCGGTCAC CCTTCTCCAG GTGGGAAGCG CTTACCTTAG GGACGCGCAA CCAGAAGGAC
                                                                        60
                                                                        100
ACGGTGAAAT TATTAAATGG GGTCTCCCTC TGGCAGGGGC
(2) INFORMATION FOR SEQ ID NO: 33:
     (i) SEQUENCE CHARACTERISTICS:
          (A) LENGTH: 100 base pairs
          (B) TYPE: nucleic acid
          (C) STRANDEDNESS: single
          (D) TOPOLOGY: linear
    (ii) MOLECULE TYPE: DNA (genomic)
   (iii) HYPOTHETICAL: NO
    (iv) ANTI-SENSE: NO
    (vi) ORIGINAL SOURCE:
          (A) ORGANISM: Homo sapiens
    (ix) FEATURE:
          (A) NAME/KEY: insertion seq
          (B) LOCATION: 14...76
          (C) IDENTIFICATION METHOD: experimental
          (D) OTHER INFORMATION:/function= "Primer for PCR cloning
                 of DNA encoding TNF-alpha analog"
                 /evidence= EXPERIMENTAL
                 /organism= "Homo sapiens"
                 /standard name= "Primer "mut30-5""
```

/label= mut30-5

/note= "Primer "mut30-5" is a synthetically synthesised 100-mer oligonucleotide comprising DNA encoding human T cell epitope P30 between stretches of DNA homologous to stretches of the human TNF-alpha gene"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 33:

GAAGGGTGAC CGATTCAACA ATTTCACCGT AAGCTTCTGG CTTCGCGTCC CTAAGGTGTC

TGCGTCGCAC CTCGAAGGGA TCATTGCCCT CTAGAGTCGA

60 100

- (2) INFORMATION FOR SEQ ID NO: 34:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..25
 - (D) OTHER INFORMATION:/label= Pep2-1
 /note= "Pep2-1 is a synthetically prepared truncated form
 of a TNF-alpha analog comprising the human T cell epitope

P2 and flanking portions of human TNF-alpha"

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 34:
- Ser Arg Thr Pro Ser Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly
 1 10 15

Ile Thr Glu Leu Gln Leu Gln Trp Leu 20 25

- (2) INFORMATION FOR SEQ ID NO: 35:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal

- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..25
 - (D) OTHER INFORMATION:/label= Pep2-3

/note= "Pep2-3 is a synthetically prepared truncated form of a TNF-alpha analog comprising the human T cell epitope P2 and flanking portions of human TNF-alpha"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 35:

Ser Gln Val Leu Phe Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly 1 5 10 15

Ile Thr Glu Leu Ile Ser Arg Ile Ala 20 25

- (2) INFORMATION FOR SEQ ID NO: 36:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..25
 - (D) OTHER INFORMATION:/label= Pep2-4
 /note= "Pep2-4 is a synthetically prepared truncated form
 of a TNF-alpha analog comprising the human T cell epitope
 P2 and flanking portions of human TNF-alpha"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 36:

Ala Glu Ala Lys Pro Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly
1 10 15

Ile Thr Glu Leu Gly Asp Arg Leu Ser 20 25

- (2) INFORMATION FOR SEQ ID NO: 37:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 25 amino acids
 - (B) TYPE: amino acid

- (C) STRANDEDNESS:
- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: peptide
- (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
- (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
- (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..25
 - (D) OTHER INFORMATION:/label= Pep2-5
 /note= "Pep2-5 is a synthetically prepared truncated form

of a TNF-alpha analog comprising the human T cell epitope P2 and flanking portions of human TNF-alpha"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 37:

Glu Lys Gly Asp Arg Gln Tyr Ile Lys Ala Asn Ser Lys Phe Ile Gly
1 5 10 15

Ile Thr Glu Leu Ser Gly Gln Val Tyr
20 25

- (2) INFORMATION FOR SEQ ID NO: 38:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..31
 - (D) OTHER INFORMATION:/label= Pep30-1

/note= "Pep30-1 is a synthetically prepared truncated form of a TNF-alpha analog comprising human T cell epitope P30 and flanking portions of human TNF-alpha"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 38:

Ser Arg Thr Pro Ser Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg 1 5 10 15

Val Pro Lys Val Ser Ala Ser His Leu Glu Arg Arg Ala Asn Ala

- (2) INFORMATION FOR SEQ ID NO: 39:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..31
 - (D) OTHER INFORMATION:/label= Pep30-2

/note= "Pep30-2 is a synthetically prepared truncated form of a TNF-alpha analog comprising the human T cell epitope P30 and flanking portions of human TNF-alpha"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 39:

Ala Leu Leu Ala Asn Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg

1 10 15

Val Pro Lys Val Ser Ala Ser His Leu Glu Gln Val Leu Phe Lys 20 25 30

- (2) INFORMATION FOR SEQ ID NO: 40:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION: 1...31
 - (D) OTHER INFORMATION:/label= Pep30-3

/note= "Pep30-3 is a synthetically prepared truncated form of a TNF-alpha analog comprising the human T cell epitope P30 and flanking portions of human TNF-alpha"

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 40:
- Tyr Ser Gln Val Leu Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg

 1 5 10 15
- Val Pro Lys Val Ser Ala Ser His Leu Glu Val Ser Tyr Gln Thr
 20 25 30
- (2) INFORMATION FOR SEQ ID NO: 41:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..31
 - (D) OTHER INFORMATION:/label= Pep30-4
 /note= "Pep30-4 is a synthetically prepared truncated
 form of a TNF-alpha analog comprising the human T cell
 epitope P30 and flanking portions of human TNF-alpha"
 - (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 41:
 - Gln Arg Glu Thr Pro Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg
 1 5 10 15
 - Val Pro Lys Val Ser Ala Ser His Leu Glu Lys Gly Asp Arg Leu
 20 25 30
- (2) INFORMATION FOR SEQ ID NO: 42:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 31 amino acids
 - (B) TYPE: amino acid
 - (C) STRANDEDNESS:
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: peptide
 - (iii) HYPOTHETICAL: NO
 - (v) FRAGMENT TYPE: internal
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens

- (ix) FEATURE:
 - (A) NAME/KEY: Peptide
 - (B) LOCATION:1..31
 - (D) OTHER INFORMATION:/label= Pep30-5

/note= "Pep30-5 is a synthetically prepared truncated form of a TNF-alpha analog comprising the human T cell epitope P30 and flanking portions of human TNF-alpha"

- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 42:
- Glu Lys Gly Asp Arg Phe Asn Asn Phe Thr Val Ser Phe Trp Leu Arg
 1 10 15

Val Pro Lys Val Ser Ala Ser His Leu Glu Gly Ile Ile Ala Leu 20 25 30

- (2) INFORMATION FOR SEQ ID NO: 43:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 1585 base pairs
 - (B) TYPE: nucleic acid
 - (C) STRANDEDNESS: double
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: cDNA
 - (iii) HYPOTHETICAL: NO
 - (iv) ANTI-SENSE: NO
 - (vi) ORIGINAL SOURCE:
 - (A) ORGANISM: Homo sapiens
 - (ix) FEATURE:
 - (A) NAME/KEY: CDS
 - (B) LOCATION:1..1585
 - (C) IDENTIFICATION METHOD: experimental
 - (D) OTHER INFORMATION:/codon_start= 1

/function= "Antigen"
/product= "TNF-alpha analog"
/evidence= EXPERIMENTAL
/gene= "tnf"
/standard name= "TNF"

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 43:

CACACCCTGA	CAAGCTGCCA	GGCAGGTTCT	CTTCCTCTCA	CATACTGACC	CACGGCTCCA	60
CCCTCTCTCC	CCTGGAAAGG	ACACCATGAG	CACTGAAAGC	ATGATCCGGG	ACGTGGAGCT	120
GGCCGAGGAG	GCGCTCCCCA	AGAAGACAGG	GGGGCCCCAG	GGCTCCAGGC	GGTGCTTGTT	180
CCTCAGCCTC	TTCTCCTTCC	TGATCGTGGC	AGGCGCCACC	ACGCTCTTCT	GCCTGCTGCA	240
CTTTGGAGTG	ATCGGCCCCC	AGAGGGAAGA	GTCCCCCAGG	GACCTCTCTC	TAATCAGCCC	300
TCTGGCCCAG	GCAGTCAGAT	CATCTTCTCG	AACCCCGAGT	GACAAGCCTG	TAGCCCATGT	360
TGTAGCAAAC	CCTCAAGCTG	AGGGGCAGCT	CCAGTGGCTG	AACCGCCGGG	CCAATGCCCT	420
CCTGGCCAAT	GGCGTGGAGC	TGAGAGATAA	CCAGCTGGTG	GTGCCATCAG	AGGGCCTGTA	480
CCTCATCTAC	TCCCAGGTCC	TCTTCAAGGG	CCAAGGCTGC	CCCTCCACCC	ATGTGCTCCT	540
CACCCACACC	ATCAGCCGCA	TCGCCGTCTC	CTACCAGACC	AAGGTCAACC	TCCTCTCTGC	600
CATCAAGAGC	CCCTGCCAGA	GGGAGACCCC	AGAGGGGGCT	GAGGCCAAGC	CCTGGTATGA	660
GCCCATCTAT	CTGGGAGGG	TCTTCCAGCT	GGAGAAGGGT	GACCGACTCA	GCGCTGAGAT	720
CAATCGGCCC	GACTATCTCG	ACTTTGCCGA	GTCTGGGCAG	GTCTACTTTG	GGATCATTGC	780

CCTGTGAGGA	GGACGAACAT	CCAACCTŤCC	CAAACGCCTC	CCCTGCCCCA	ATCCCTTTAT	840
TACCCCCTCC	TTCAGACACC	CTCAACCTCT	TCTGGCTCAA	AAAGAGAATT	GGGGGCTTAG	900
GGTCGGAACC	CAAGCTTAGA	ACTTTAAGCA	ACAAGACCAC	CACTTCGAAA	CCTGGGATTC	960
AGGAATGTGT	GGCCTGCACA	GTGAAGTGCT	GGCAACCACT	AAGAATTCAA	ACTGGGGCCT	1020
CCAGAACTCA	CTGGGGCCTA	CAGCTTTGAT	CCCTGACATC	TGGAATCTGG	AGACCAGGGA	1080
GCCTTTGGTT	CTGGCCAGAA	TGCTGCAGGA	CTTGAGAAGA	CCTCACCTAG	AAATTGACAC	1140
AAGTGGACCT	TAGGCCTTCC	TCTCTCCAGA	TGTTTCCAGA	CTTCCTTGAG	ACACGGAGCC	1200
CAGCCCTCCC	CATGGAGCCA	GCTCCCTCTA	TTTATGTTTG	CACTTGTGAT	TATTTATTAT	1260
TTATTTATTA	TTTATTTATT	TACAGATGAA	TGTATTTATT	TGGGAGACCG	GGGTATCCTG	1320
GGGGACCCAA	TGTAGGAGCT	GCCTTGGCTC	AGACATGTTT	TCCGTGAAAA	CGGAGGCTGA	1380
ACAATAGGCT	GTTCCCATGT	AGCCCCCTGG	CCTCTGTGCC	TTCTTTTGAT	TATGTTTTTT	1440
AAAATATTAT	CTGATTAAGT	TGTCTAAACA	ATGCTGATTT	GGTGACCAAC	TGTCACTCAT	1500
TGCTGAGGCC	TCTGCTCCCC	AGGGAGTTGT	GTCTGTAATC	GGCCTACTAT	TCAGTGGCGA	1560
GAAATAAAGG	TTGCTTAGGA	AAGAA				1585

(2) INFORMATION FOR SEQ ID NO: 44:

- (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH: 233 amino acids
 - (B) TYPE: amino acid
 - (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: protein
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 44:

Met Ser Thr Glu Ser Met Ile Arg Asp Val Glu Leu Ala Glu Glu Ala 1 5 10 15

Leu Pro Lys Lys Thr Gly Gly Pro Gln Gly Ser Arg Arg Cys Leu Phe
20 25 30

Leu Ser Leu Phe Ser Phe Leu Ile Val Ala Gly Ala Thr Thr Leu Phe 35 40 45

Cys Leu Leu His Phe Gly Val Ile Gly Pro Gln Arg Glu Glu Phe Pro 50 55 60

Arg Asp Leu Ser Leu Ile Ser Pro Leu Ala Gln Ala Val Arg Ser Ser 65 70 75 80

Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His Val Val Ala Asn Pro 85 90 95

Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg Arg Ala Asn Ala Leu 100 105 110

Leu Ala Asn Gly Val Glu Leu Arg Asp Asn Gln Leu Val Val Pro Ser 115 120 125

Glu Gly Leu Tyr Leu Ile Tyr Ser Gln Val Leu Phe Lys Gly Gln Gly
130 135 140

Cys Pro Ser Thr His Val Leu Leu Thr His Thr Ile Ser Arg Ile Ala 145 150 155 160

Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser Ala Ile Lys Ser Pro 165 170 175

Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala Lys Pro Trp Tyr Glu 180 185 190

Pro Ile Tyr Leu Gly Gly Val Phe Gln Leu Glu Lys Gly Asp Arg Leu 195 200 205

Ser Ala Glu Ile Asn Arg Pro Asp Tyr Leu Asp Phe Ala Glu Ser Gly 210 215 220

Gln Val Tyr Phe Gly Ile Ile Ala Leu 225 230